## Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application:

## Listing of Claims:

1. (Currently amended) A compound of the Formula Ia:

or a pharmaceutically acceptable salt thereof, wherein.

L- is a Ligand unit;

-A- is a Stretcher unit;

a is 1;

each -W- is independently an Amino Acid unit;

-Y- is a self-immolative Spacer unit;

w is an integer ranging from 2 to 12;

y is 1 or 2;

p ranges from 1 to about 20; and

-D is a Drug unit of the formula:

wherein, the wavy line indicates the point of attachment to the Spacer unit, and

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## independently at each location:

R2 is selected from the group consisting of -H and -C1-C8 alkyl;

 $R^3$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle);

R<sup>4</sup> is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle) wherein; R<sup>5</sup> is selected from the group consisting of -H and -methyl; or R<sup>4</sup> and R<sup>5</sup> join and form a ring with the carbon atom to which they are attached and R<sup>4</sup> and R<sup>5</sup> have the formula -(CR\*R<sup>b</sup>)<sub>n</sub>- wherein; R<sup>a</sup> and R<sup>b</sup> are independently selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl and -C<sub>3</sub>-C<sub>8</sub> carbocycle and n is selected from the group consisting of 2, 3, 4, 5 and 6:

R6 is selected from the group consisting of -H and -C1-C8 alkyl;

 $R^7$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle):

each  $R^8$  is independently selected from the group consisting of -H, -OH, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle and -O-(C<sub>1</sub>-C<sub>8</sub> alkyl);

R<sup>9</sup> is selected from the group consisting of -H and -C<sub>1</sub>-C<sub>8</sub> alkyl; R<sup>10</sup> is selected from the group consisting of:

Z is -O-, -S-,-NH- or -N(R<sup>14</sup>)-;

 $R^{11}$  is selected from the group consisting of -H, -OH, -NH<sub>2</sub>, -NHR<sup>14</sup>, -N( $R^{14}$ )<sub>2</sub>, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>2</sub>-C<sub>8</sub> carbocycle, -O<sub>-</sub>(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>2</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle); or  $R^{11}$  is an oxygen atom which forms a carbonyl unit (C=O) with the carbon atom to which it is attached and a hydrogen atom on this carbon atom is replaced by one of the bonds in the (C=O) double bond;

each  $R^{12}$  is independently selected from the group consisting of -aryl and -C<sub>3</sub>-C<sub>8</sub> heterocycle;

 $R^{13} is selected from the group consisting of -H, -OH, -NH2, -NHR^{14}, -N(R^{14})_2, -C_1-C_8 alkyl, -C_3-C_8 carbocycle, -O-(C_1-C_8 alkyl), -aryl, -C_1-C_8 alkyl-aryl, -C_1-C_8 alkyl-(C_3-C_8 carbocycle), C_3-C_8 heterocycle and -C_{1-8} alkyl-(C_3-C_8 heterocycle); and -C_{1-8} alkyl-(C_3-C_8 heterocycle); and$ 

each R14 is independently -H or -C1-C8 alkyl.

2-6. (Canceled)

7. (Currently amended) A compound of the formula Ia:

$$L - \left( -A_{\overline{a}} - W_{\overline{w}} - Y_{\overline{y}} - D \right)_{p}$$

or a pharmaceutically acceptable salt thereof, wherein,

L- is a Ligand unit;

-A- is a Stretcher unit;

a is 1:

each -W- is independently an Amino Acid unit;

-Y- is a self-immolative Spacer unit;

w is an integer ranging from 2 to 12;

v is 1 or 2:

p ranges from 1 to about 20; and

-D is a Drug unit having the structure:

or a pharmaceutically acceptable salt thereof,

wherein, the wavy line [[is]] indicates the point of attachment to the Spacer unit, and independently at each location:

R2 is selected from the group consisting of -H and -methyl;

R<sup>3</sup> is selected from the group consisting of -H, -methyl, and -isopropyl;

R4 is selected from the group consisting of -H and -methyl;

R<sup>5</sup> is selected from the group consisting of -isopropyl, -isobutyl, -sec-butyl, -

methyl and -t-butyl or  $R^4$  and  $R^5$  join[[,]] and form a ring with the carbon atom to which they are attached and  $R^4$  and  $R^5$  have the formula -( $CR^aR^b$ )<sub>n</sub>- where<u>in</u>:  $R^a$  and  $R^b$  are independently selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, and

-C<sub>3</sub>-C<sub>8</sub> carbocycle, and n is selected from the group consisting of 2, 3, 4, 5 and 6;

R<sup>6</sup> is selected from the group consisting of -H and -methyl;

each R8 is independently selected from the group consisting of -OH, -methoxy

and -ethoxy;

R10 is selected from the group consisting of:

$$\mathbb{R}^{24}$$
O and  $\mathbb{R}^{27}$ CH<sub>3</sub>

 $R^{24}$  is selected from the group consisting of H and -C(O) $R^{25}$ -; wherein  $R^{25}$  is selected from the group consisting of -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>2</sub>-C<sub>8</sub> carbocycle, -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>2</sub>-C<sub>8</sub> carbocycle), -C<sub>2</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>2</sub>-C<sub>8</sub> heterocycle);

 $Z \text{ is -O-, -NH-, -OC(O)-, -NHC(O)-, or -NR}^{28}C(O)-; \\ \text{ where } R^{28} \text{ is selected from the group consisting of -H and -C}_1-C_8 \text{ alkyl}; }$ 

n is 0 or 1; and

 $R^{27} is selected from the group consisting of -H, -N_3, -C_1-C_8 alkyl, -C_3-C_8 carbocycle, -aryl, -C_1-C_8 alkyl-aryl, -C_1-C_8 alkyl-(C_3-C_8 carbocycle), -C_3-C_8 heterocycle and -C_1-C_8 alkyl-(C_3-C_8 heterocycle) when n is 0; and <math>R^{27}$  is selected from the group consisting of -H, -C\_1-C\_8 alkyl, -C\_3-C\_8 carbocycle, -aryl, -C\_1-C\_8 alkyl-aryl, -C\_1-C\_8 alkyl-(C\_3-C\_8 carbocycle), -C\_3-C\_8 heterocycle and -C\_1-C\_8 alkyl-(C\_3-C\_8 heterocycle) when n is 1.

- (Canceled)
- (Currently amended) [[A]]The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -D is a Drug unit having the structure:

10-16. (Canceled)

- (Currently amended) [[A]]The compound or a pharmaceutically acceptable salt of the compound of claim 1 or claim 7 wherein the Ligand unit is an antibody.
- (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 17 wherein the antibody is a monoclonal antibody.
- 19. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 18 wherein the monoclonal antibody specifically binds the CD30 antigen, the CD20 antigen, the Lewis X or Y antigen, the CD33 antigen, the CD38 antigen, the CD19 antigen, the CA15-3 antigen or the epidermal growth factor antigen.
- (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -Yy- is;

Q is selected from the group consisting of  $-C_1-C_8$  alkyl,  $-O-(C_1-C_8$  alkyl), -halogen, -nitro and -evano; and

m is an integer ranging from 0-4, the amino terminus of -Yy- forming a bond with the Amino acid unit and the other terminus of -Yy- forming a bond with the Drug unit.

21. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is;

and r is an integer ranging from 1-10, the carbonyl terminus of -A- forming a bond with the Amino Acid unit and the succinimido terminus of -A- forming a bond with the Ligand unit.

22. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is:

and r is an integer ranging from 1-10, the carbonyl terminus of -A- forming a bond with the Amino Acid unit and the amidomethyl terminus of -A- forming a bond with the Ligand unit.

 (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is;

and r is an integer ranging from 1-10, the carbonyl terminus of -A- forming a bond with the Amino acid unit and the succinimido terminus of -A- forming a bond with the Ligand unit.

24. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is:

$$\begin{array}{c} \underbrace{ \left\{ \begin{array}{c} \\ \\ \\ \end{array} \right\} } \\ \text{N-(CH}_2\text{CH}_2\text{O)}_1\text{CH}_2\text{C(O)} \\ -\underbrace{ \left\{ \begin{array}{c} \\ \\ \end{array} \right\} } \\ \end{array}$$

and r is an integer ranging from 1-10, the carbonyl terminus of -A- forming a bond with the Amino acid unit and the succinimido terminus of -A- forming a bond with the Ligand unit.

25. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is:

$$e^{\int_{\mathbb{R}^{2}}^{\mathbb{R}^{2}} \left( \int_{\mathbb{R}^{2}}^{\mathbb{R}^{2}} \int_{\mathbb{R}^{2}}^{\mathbb{R}^{2}$$

and r is an integer ranging from 1-10, the carbonyl terminus of -A- forming a bond with the Amino acid unit and the amidomethyl terminus of -A- forming a bond with the Ligand unit.

 (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -A- is;

the carbonyl terminus of -A- forming a bond with the Amino acid unit and the amidomethyl terminus of -A- forming a bond with the Ligand unit.

 (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 21 wherein -A- is;

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the carbonyl terminus of -A- forming a bond with the Amino acid unit and the succinimido terminus of -A- forming a bond with the Ligand unit.

28. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 22 wherein -A- is:

the carbonyl terminus of -A- forming a bond with the Amino acid unit and the amidomethyl terminus of -A- forming a bond with the Ligand unit.

29. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 24 wherein -A- is:

the carbonyl terminus of -A- forming a bond with the Amino acid unit and the succinimido terminus of -A- forming a bond with the Ligand unit.

- 30. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -Ww- is -Phenylalanine-Lysine-, the amino terminus of -Ww- forming a bond with the Stretcher unit and the C- terminus of -Ww-forming a bond with the Spacer unit.
  - 31-43. (Canceled)
  - 44. (Currently amended) A compound of the formula:

 $R^{16} \xrightarrow[R^2]{} N \xrightarrow[N^2]{} N \xrightarrow[N^4]{} R^5 \xrightarrow[R^6]{} R^8 \xrightarrow[N^8]{} 0 \xrightarrow[R^8]{} N \xrightarrow[N^8]{} R^{11}$ 

or a pharmaceutically acceptable salt thereof; wherein, independently at each location:

R2 is selected from the group consisting of -H and -C1-C8 alkyl;

R3 is selected from the group consisting of -H, -C1-C8 alkyl, -C3-C8

carbocycle,  $-O-(C_1-C_8 \text{ alkoy})$ , -aryl,  $-C_1-C_8 \text{ alkyl-aryl}$ ,  $-C_1-C_8 \text{ alkyl-(}C_3-C_8 \text{ carbocycle)}$ ,  $-C_3-C_8 \text{ heterocycle}$  and  $-C_1-C_8 \text{ alkyl-(}C_3-C_8 \text{ heterocycle)}$ ;

 $R^4$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub>

carbocycle, -O- $(C_1-C_8 \text{ alkvy})$ , -aryl,  $-C_1-C_8 \text{ alkyl-aryl}$ ,  $-C_1-C_8 \text{ alkyl-}(C_3-C_8 \text{ carbocycle})$ ,  $-C_3-C_8$  heterocycle and  $-C_1-C_8 \text{ alkyl-}(C_3-C_8 \text{ heterocycle})$  wherein;  $R^5$  is selected from the group consisting of -H and -methyl; or  $R^4$  and  $R^5$  join and form a ring with the carbon atom to which they are attached and  $R^4$  and  $R^5$  have the formula;  $-(C_1R^3R^6)_n$ - wherein;  $R^a$  and  $R^b$  are independently selected from the group consisting of -H,  $-C_1-C_8$  alkyl and  $-C_3-C_8$  carbocycle and  $R^4$  is selected from the group consisting of 2, 3, 4, 5 and 6:

R<sup>6</sup> is selected from the group consisting of -H and -C<sub>1</sub>-C<sub>8</sub> alkyl;

R7 is selected from the group consisting of -H, -C1-C8 alkyl, -C3-C8

carbocycle,  $-O-(C_1-C_8 \text{ alkyx})$ , -aryl,  $-C_1-C_8 \text{ alkyl-aryl}$ ,  $-C_1-C_8 \text{ alkyl-(C_3-C_8 \text{ carbocycle)}}$ ,  $-C_3-C_8 \text{ heterocycle}$  and  $-C_1-C_8 \text{ alkyl-(C_3-C_8 \text{ heterocycle)}}$ ;

each  $R^8$  is independently selected from the group consisting of -H, -OH, -  $C_1$ - $C_8$  alkvl, - $C_3$ - $C_8$  carbocycle and -O-( $C_1$ - $C_8$  alkvxy);

 $R^9$  is selected from the group consisting of -H and -C1-C8 alkyl;

R<sup>11</sup> is selected from the group consisting of -H, -OH, -NH<sub>2</sub>, -NHR<sup>14</sup>, -

 $N(R^{14})_2, -C_1 - C_8 \text{ alkyl}, -C_3 - C_8 \text{ carbocycle, -O-}(C_1 - C_8 \text{ alkyl}), -\text{aryl, -C}_1 - C_8 \text{ alkyl-aryl, -C}$ 

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(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle); or R<sup>11</sup> is an oxygen atom which forms a carbonyl unit (C=O) with the carbon atom to which it is attached and a hydrogen atom on this carbon atom is replaced by one of the bonds in the (C=O) double bond; each R<sup>12</sup> is independently selected from the group consisting of -aryl and -

## C3-C8 heterocycle;

 $R^{13} \ is \ selected \ from \ the \ group \ consisting \ of -H, -OH, -NH2, -NHR^{14}, -N(R^{14})_2, -C_1-C_8 \ alkyl, -C_2-C_8 \ carbocycle, -O-(C_1-C_8 \ alkoxy), -aryl, -C_1-C_8 \ alkyl-aryl, -C_1-C_8 \ alkyl-(C_3-C_8 \ carbocycle), -C_3-C_8 \ heterocycle \ and -C_1-C_8 \ alkyl-(C_3-C_8 \ heterocycle);$ 

# R<sup>16</sup> is A'a-Ww-Yy-

#### wherein

each -W- is independently an Amino Acid unit;

-Y- is a self-immolative Spacer unit;

w is an integer ranging from 2 to 12;

y is 1 or 2;

-A' is a Stretcher unit; and

a is 1.

45. (Currently amended) The compound of claim 44 having the structure:

$$\begin{array}{c} \begin{array}{c} \begin{array}{c} \\ \\ \\ \\ \\ \\ \\ \\ \end{array} \end{array}$$

46. (Currently amended) The compound of claim 44 having the structure:

or a pharmaceutically acceptable salt thereof.

- 47. (Canceled)
- 48. (Currently amended) The compound of claim 44 having the structure:

- 49-51. (Canceled)
- 52. (Currently amended) The compound of claim 44 having the structure:

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or a pharmaceutically acceptable salt thereof.

- 53. (Canceled)
- 54. (Currently amended) The compound of claim 128 having the structure:

- 55. (Canceled)
- 56. (Currently amended) The compound of claim 1 having the structure:

or a pharmaceutically acceptable salt thereof.

- 57-58. (Canceled)
- 59. (Currently amended) The compound of claim 1 having the structure:

- 60-76. (Canceled)
- 77. (Currently amended) The compound of claim 1 having the formula:

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or a pharmaceutically acceptable salt thereof, wherein L is a monoclonal antibody.

- 78. (Canceled)
- (Previously presented) The compound of claim 54 or a pharmaceutically 79. acceptable salt thereof, wherein L is a monoclonal antibody.
  - 80-99. (Canceled)
- 100. (Previously presented) The compound or pharmaceutically acceptable salt thereof of claim 79 wherein L specifically binds the CD20 antigen.
  - 101-103. (Canceled)
- 104. (Previously presented) The compound or pharmaceutically acceptable salt thereof of claim 77 wherein L specifically binds the CD20 antigen.
  - 105-110. (Canceled)
- (Previously presented) A composition comprising an effective amount of a compound or a pharmaceutically acceptable salt thereof of claim 1 or claim 7, and a pharmaceutically acceptable carrier or vehicle.
  - 112-118. (Canceled)
- 119. (Previously presented) The compound or a pharmaceutically acceptable salt thereof of claim 1 in an isolated or a purified form.
  - 120. (Canceled)

- 121. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein -Ww- is -valine-citrulline-, the amino terminus of -Ww- forming a bond with the Stretcher unit, and the C- terminus of -Ww- forming a bond with a the Spacer unit.
- (Currently amended) The compound of claim 44 or a pharmaceutically acceptable salt of the compound of claim 44, wherein

-A' is selected from the group consisting of:

## wherein

G is selected from the group consisting of -Cl, -Br, -I, -O-mesyl and -O-tosyl;

J is selected from the group consisting of -Cl, -Br, -I, -F, -OH, -O-N-succinimide, -O-(4-nitrophenyl), -O-pentafluorophenyl, -O-tetrafluorophenyl and -O-C(O)-OR<sup>18</sup>;

ffa is 1:11

 $R^{17}$  is selected from the group consisting of  $-C_1-C_{10}$  alkylene-,  $-C_3-C_8$  carbocyclo-,  $-O_1-C_{10}$  alkylene-,  $-C_1-C_{10}$  alkylene-,  $-C_1-C_{10}$  alkylene-,  $-C_1-C_{10}$  alkylene-,  $-C_1-C_{10}$  alkylene-,  $-C_1-C_{10}$  alkylene-,  $-C_1-C_1$ 0 alkylene-,  $-C_2-C_8$  heterocyclo-,  $-C_1-C_1$ 0 alkylene-,  $-C_3-C_8$  heterocyclo-,  $-C_1-C_1$ 0 alkylene-,  $-C_3-C_8$  heterocyclo-,  $-C_1-C_1$ 0 alkylene-,  $-C_1-C_1$ 0 alkylen

r is an integer ranging from 1-10; and

R18 is -C1-C8 alkyl or -aryl.

- 123. (Canceled)
- 124. (Previously presented) A composition comprising an effective amount of a compound or a pharmaceutically acceptable salt thereof of claim 79 and a pharmaceutically acceptable carrier or vehicle.
- 125. (Previously presented) A composition comprising an effective amount of a compound or a pharmaceutically acceptable salt thereof of claim 121 and a pharmaceutically acceptable carrier or vehicle.
- 126. (Previously presented) The compound or a pharmaceutically acceptable salt thereof of claim 79 in an isolated or a purified form.
- 127. (Previously presented) The compound or a pharmaceutically acceptable salt thereof of claim 121 in an isolated or a purified form.
- 128. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein

-Aa-Ww-Yy- has the formula:

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the succinimido terminus forming a bond with the Ligand unit and the other terminus forming a bond with the Drug unit.

129. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claim 7 wherein

-Aa-Ww-Yy- has the formula:

the succinimido terminus forming a bond with the Ligand unit and the other terminus forming a bond with the Drug unit.

- 130. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claims 128 or 129 wherein the ligand unit is a monoclonal antibody.
- 131. (Currently amended) The compound or pharmaceutically acceptable salt thereof of claim 1 wherein R10 is

132. (Currently amended) The compound or pharmaceutically acceptable salt thereof of claim 7 wherein  $R^{10}$  is:

- 133. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claim 19 wherein the monoclonal antibody specifically binds the CD30 antigen.
- 134. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claim 19 wherein the monoclonal antibody specifically binds the CD19 antigen.
- 135. (Previously presented) The compound or a pharmaceutically acceptable salt of the compound of claim 19 wherein the monoclonal antibody specifically binds the CD33 antigen.
- 136. (Currently amended) The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein  $-A_a$  is:

 $\label{eq:wherein} \begin{array}{l} wherein \ R^{17} \ is \ selected from \ the group \ consisting \ of \ -C_1-C_{10} \ alkylene, \ C_3-C_8 \ carbocyclo-, \ -O-(C_1-C_8 \ alkyl)-, \ -arylene-, \ -C_1-C_{10} \ alkylene-, \ -arylene-C_1-C_{10} \ alkylene-, \ -C_1-C_{10} \ alkylene-, \ -C_1-C_10 \ alkylene-,$ 

- 137. (Currently amended) [[A]]The compound or a pharmaceutically acceptable salt of the compound of claim 1 wherein p ranges from 1 to about 5.
- 138. (Currently amended) [[A]]<u>The</u> compound or a pharmaceutically acceptable salt of the compound of claim 79 wherein p ranges from 1 to about 5.
- 139. (Currently amended) [[A]]<u>The</u> compound or a pharmaceutically acceptable salt of the compound of claim 54 where<u>in</u> L is a monoclonal antibody that specifically binds the CD30 antigen, the CD20 antigen, the Lewis X or Y antigen, the CD33 antigen, the CD19 antigen, the CD38 antigen, the CEA antigen, the CA15-3 antigen or the epidermal growth factor antigen.
- 140. (Currently amended) [[A]]The compound or a pharmaceutically acceptable salt of the compound of claim 139 wherein the monoclonal antibody specifically binds the CD30 antigen.
- 141. (Currently amended) A composition comprising drug-linker-ligand conjugates having Formula Ia:

$$L - \left( A_{\overline{a}} - W_{\overline{w}} - Y_{\overline{y}} - D \right)_{p}$$

or a pharmaceutically acceptable salt thereof; wherein.

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L- is a Ligand unit;

-A- is a Stretcher unit:

a is 1:

each -W- is independently an Amino Acid unit;

-Y- is a self-immolative Spacer unit;

w is an integer ranging from 2 to 12;

y is 1 or 2;

p ranges from 1 to about 5 and is the average number of -A\_a-W\_w-Y\_y-D units per ligand in the composition; and

-D is a Drug unit of the formula:

wherein, the wavy line indicates the point of attachment to the Spacer unit, and independently at each location:

 $R^2$  is selected from the group consisting of -H and -C1-C8 alkyl;

 $R^2$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O<sub>-</sub>(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle);

 $R^4$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle) wherein;  $R^5$  is selected from the group consisting of -H and -methyl; or  $R^4$  and  $R^5$  join and form a ring with the carbon atom to which they are attached and

R<sup>4</sup> and R<sup>5</sup> have the formula -(CR<sup>a</sup>R<sup>b</sup>)<sub>n</sub>, wherein; R<sup>a</sup> and R<sup>b</sup> are independently selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl and -C<sub>3</sub>-C<sub>8</sub> carbocycle and n is selected from the group consisting of 2, 3, 4, 5 and 6;

R6 is selected from the group consisting of -H and -C1-C8 alkyl;

 $R^7$  is selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>9</sub> alkyl), -aryl.

 $-C_1-C_8 \ alkyl-aryl, -C_1-C_8 \ alkyl-(C_3-C_8 \ carbocycle), -C_3-C_8 \ heterocycle \ and -C_1-C_8 \ alkyl-(C_3-C_8 \ heterocycle);$ 

each  $R^8$  is independently selected from the group consisting of -H, -OH, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle and -O-(C<sub>1</sub>-C<sub>8</sub> alkyl);

R9 is selected from the group consisting of -H and -C1-C8 alkyl;

R<sup>10</sup> is selected from the group consisting of:

Z is -O-, -S-,-NH- or -N( $\mathbb{R}^{14}$ )-;

 $R^{11}$  is selected from the group consisting of -H, -OH, -NH<sub>2</sub>, -NHR<sup>14</sup>, -N(R<sup>14</sup>)<sub>2</sub>, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -O-(C<sub>1</sub>-C<sub>8</sub> alkyl), -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle); or  $R^{11}$  is an oxygen atom which forms a carbonyl unit (C=O) with the carbon atom to which it is attached and a hydrogen atom on this carbon atom is replaced by one of the bonds in the (C=O) double bond;

each R<sup>12</sup> is independently selected from the group consisting of -aryl and -C<sub>3</sub>-C<sub>8</sub>

heterocycle;

 $R^{13} \ is \ selected from the group consisting of -H, -OH, -NH_2, -NHR^{14}, -N(R^{14})_2, -C_1-C_8 \ alkyl, -C_3-C_8 \ carbocycle, -O-(C_1-C_8 \ alkyl), -aryl, -C_1-C_8 \ alkyl-aryl, -C_1-C_8 \ alkyl-(C_3-C_8 \ carbocycle), C_3-C_8 \ heterocycle \ and -C_{1-8} \ alkyl-(C_3-C_8 \ heterocycle); \ and$ 

each  $R^{14}$  is independently -H or -C<sub>1</sub>-C<sub>8</sub> alkyl.

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142. (Currently amended) A composition comprising drug-linker-ligand conjugates having Formula Ia:

$$L - \left( -A_{\overline{a}} - W_{\overline{w}} - Y_{\overline{y}} - D \right)_{p}$$
Ia

or a pharmaceutically acceptable salt thereof wherein,

L- is a Ligand unit;

-A- is a Stretcher unit:

a is 1;

each -W- is independently an Amino Acid unit;

-Y- is a self-immolative Spacer unit;

w is an integer ranging from 2 to 12;

y is 1 or 2;

p ranges from 1 to about 5 and is the average number of  $-A_{a^*}W_{w^*}Y_{y^*}D$  units per ligand in the composition; and

-D is a Drug unit having the structure:

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## or a pharmaceutically acceptable salt thereof,

wherein, the wavy line [[is]]indicates the point of attachment to the Spacer unit, and independently at each location:

R<sup>2</sup> is selected from the group consisting of -H and -methyl;

R<sup>3</sup> is selected from the group consisting of -H, -methyl, and -isopropyl;

R4 is selected from the group consisting of -H and -methyl;

R<sup>5</sup> is selected from the group consisting of -isopropyl, -isobutyl, -sec-butyl, -

methyl and -t-butyl or  $R^4$  and  $R^5$  join[[,]] and form a ring with the carbon atom to which they are attached and  $R^4$  and  $R^5$  have the formula -( $CR^aR^b$ )<sub>n</sub>- where<u>in:</u>  $R^a$  and  $R^b$  are independently selected from the group consisting of -H, -C<sub>1</sub>-C<sub>8</sub> alkyl, and

-C<sub>3</sub>-C<sub>8</sub> carbocycle, and n is selected from the group consisting of 2, 3, 4, 5 and 6;

 $\ensuremath{R^6}$  is selected from the group consisting of -H and -methyl;

each R<sup>8</sup> is independently selected from the group consisting of -OH, -methoxy

and -ethoxy;

R<sup>10</sup> is selected from the group consisting of:

$$\mathbb{R}^{24}$$
O and  $\mathbb{R}^{27}$  $\mathbb{R$ 

 $R^{24} \ is \ selected \ from \ the \ group \ consisting \ of \ H \ and \ -C(O)R^{25}-; \ wherein; \ R^{25} \ is \ selected \ from \ the \ group \ consisting \ of \ -C_1-C_8 \ alkyl, \ -C_3-C_8 \ carbocycle, \ -aryl, \ -C_1-C_8 \ alkyl-aryl, \ -C_1-C_8 \ alkyl-(C_3-C_8 \ carbocycle), \ -C_3-C_8 \ heterocycle \ and \ -C_1-C_8 \ alkyl-(C_3-C_8 \ heterocycle);$ 

Z is -O-, -NH-, -OC(O)-, -NHC(O)-, or -NR<sup>28</sup>C(O)-; where; R<sup>28</sup> is selected from

the group consisting of -H and -C1-C8 alkyl;

n is 0 or 1; and

 $R^{27}$  is selected from the group consisting of -H<sub>1</sub>-N<sub>3</sub>, -C<sub>1</sub>-C<sub>8</sub> alkyl, -C<sub>3</sub>-C<sub>8</sub> carbocycle, -aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-aryl, -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> carbocycle), -C<sub>3</sub>-C<sub>8</sub> heterocycle and -C<sub>1</sub>-C<sub>8</sub> alkyl-(C<sub>3</sub>-C<sub>8</sub> heterocycle) when n is 0; and

 $R^{27} \ is \ selected \ from \ the \ group \ consisting \ of \ -H, \ -C_1-C_8 \ alkyl, \ -C_3-C_8 \ carbocycle, \ -aryl, \ -C_1-C_8 \ alkyl-aryl,$ 

 $-C_1-C_8 \ alkyl-(C_3-C_8 \ carbocycle), -C_3-C_8 \ heterocycle \ and -C_1-C_8 \ alkyl-(C_3-C_8 \ heterocycle) \ when \ n \ is \ l.$ 

143. (Currently amended) The composition of claim 141 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, R<sup>10</sup> is

144. (Currently amended) The composition of claim 142 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, R<sup>10</sup> is

145. (Currently amended) The composition of claim 141 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof. -D is a Drug unit having the structure:

or a pharmaceutically acceptable salt thereof.

146. (Currently amended) The composition of claim 141 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, -Aa-Ww-Yy- has the formula:

the succinimido terminus forming a bond with the Ligand unit and the other terminus forming a bond with the Drug unit.

147. (Currently amended) The composition of claim 142 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, -Aa-Ww-Yy- has the formula:

the succinimido terminus forming a bond with the Ligand unit and the other terminus forming a bond with the Drug unit.

- 148. (Currently amended) The composition of claim 141 where<u>in in the drug-linker-ligand conjugates or pharmaceutically acceptable salt thereof</u>, the ligand unit is a monoclonal antibody.
- 149. (Currently amended) The composition of claim 148 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD30 antigen, the CD20 antigen, the CD19 antigen, the Lewis X or Y antigen, the CD33 antigen, the CD38 antigen, the CEA antigen, the CA15-3 antigen or the epidermal growth factor antigen.
- 150. (Currently amended) The composition of 149 wherein in the drug-linkerligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD19 antigen.
- 151. (Currently amended) The composition of claim 149 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD30 antigen.
- 152. (Currently amended) The composition of claim 149 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD33 antigen.

153. (Currently amended) The composition of claim 147 wherein the druglinker-ligand conjugates have the formula:

- 154. (Currently amended) The composition of claim 153 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, L is a monoclonal antibody.
- 155. (Currently amended) The composition of claim 154 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD20 antigen, the CD30 antigen, the CD33 antigen, the CD19 antigen, the CD38 antigen, the CA15-3 antigen, the CEA antigen, or the epidermal growth factor antigen.
- 156. (Currently amended) The composition of claim 155 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the the CD30 antigen.
- 157. (Currently amended) The composition of claim 155 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD19 antigen.
- 158. (Currently amended) The composition of claim 155 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD20 antigen.

- 159. (Currently amended) The composition of claim 155 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD33 antigen.
- 160. (Currently amended) The composition of claim 142 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof. L is a monoclonal antibody.
- 161. (Currently amended) The composition of claim 160 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD20 antigen, the CD30 antigen, the CD33 antigen, the CD19 antigen, the CD38 antigen, the CA15-3 antigen, the CEA antigen, or the epidermal growth factor antigen.
- 162. (Currently amended) The composition of claim 161 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the monoclonal antibody specifically binds the CD30 antigen.
- 163. (Currently amended) The composition of claim 154 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, the antibody is attached to the drug mojety through a cysteine residue of the antibody.
- 164. (Currently amended) The compound of claim 122 or a pharmaceutically acceptable salt of the compound of claim 122, wherein

Aa- is:

 $\label{eq:wherein} $$ R^{17}$ is selected from the group consisting of $-C_1-C_{10}$ alkylene, $C_3-C_8$ carbocyclo-, $-O_1(C_1-C_8$ alkyl)-, $-arylene-, $-C_1-C_{10}$ alkylene-arylene-, $-arylene-C_1-C_{10}$ alkylene-, $-C_1-C_{10}$ alkylene-, $-C_3-C_8$ carbocyclo-, $-(C_3-C_8$ carbocyclo-)-C_1-C_{10}$ alkylene-, $-C_3-C_8$ heterocyclo-$ 

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- , -C<sub>1</sub>-C<sub>10</sub> alkylene-(C<sub>3</sub>-C<sub>8</sub> heterocyclo)-, (C<sub>3</sub>-C<sub>8</sub> heterocyclo)-C<sub>1</sub>-C<sub>10</sub>alkylene-, -(CH<sub>2</sub>CH<sub>2</sub>O)<sub>r</sub>-, and -(CH<sub>2</sub>CH<sub>2</sub>O)<sub>r</sub>-CH<sub>2</sub>-; and r is an integer ranging from 1-10.
- $165. \quad \text{(Currently amended) The compound } \frac{\text{of elaim-1}}{\text{or a pharmaceutically acceptable salt of the compound } \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically acceptable salt of the compound }} \frac{1}{\text{or a pharmaceutically }} \frac{1}{$
- $166. \quad \text{(Currently amended) The composition of claim 141 wherein $\underline{in$ the drug-linker-ligand conjugates or pharmaccutically acceptable salt thereof, $R^2$ is $-C_1-C_8$ alkyl.}$
- 167. (Currently amended) The compound of elaim 7 or a pharmaceutically acceptable salt of the compound of claim 7 wherein  $\mathbb{R}^2$  is -methyl.
- 168. (Currently amended) The composition of claim 142 wherein in the druglinker-ligand conjugates or pharmaceutically acceptable salt thereof, R<sup>2</sup> is -methyl.